

# Are Australian children iodine deficient?

## Results of the Australian National Iodine Nutrition Study

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**E**nvironmental iodine deficiency causes a wide spectrum of devastating mental and physical disorders, collectively described as iodine deficiency disorders.<sup>1</sup> While endemic goitre is the most visible consequence of iodine deficiency, the most significant and profound consequences are on the developing brain.<sup>2</sup> Impaired intellectual development of people living in iodine-deficient regions is of particular concern, especially when all of the adverse effects of iodine deficiency can be prevented by long-term, sustainable iodine prophylaxis.

For over two decades the Australian Centre for Control of Iodine Deficiency Disorders (ACCIDD) has performed sporadic surveys of urinary iodine excretion (UIE) levels in Australia. In 1992, we reported that the median UIE levels in the Australian population were above 200 µg/L, consistent with iodine sufficiency.<sup>3</sup> However, this situation has changed dramatically in recent years, with several studies from Victoria, New South Wales<sup>4-8</sup> and Tasmania,<sup>9,10</sup> showing median UIE levels of <100 µg/L. In pregnant women, median UIE levels have been found to be well below 100 µg/L, indicating that around half of these women in NSW, Victoria and Tasmania are iodine deficient.<sup>4,5</sup>

The decline in iodine intake in Australia appears to be due to changes within the dairy industry, with chlorine-containing sanitisers now replacing iodine-containing sanitisers. Iodine released from these chemicals into milk has been the major source of dietary iodine in Australia for at least four decades, but is now declining. Another contributory factor to the re-emergence of iodine deficiency in Australia has been the decreasing consumption of iodised salt.<sup>3</sup> To our knowledge, few if any food manufacturers use iodised salt in the preparation and manufacture of foods.

Our study was conducted with the aim of documenting the population iodine nutritional status of schoolchildren in Australia.

### METHODS

#### Approval and consent

The study was approved by the Western Sydney Area Health Service Human

### ABSTRACT

**Objective:** To document the population iodine nutritional status in Australian schoolchildren.

**Design and setting:** Cross-sectional survey of schoolchildren aged 8–10 years, based on a one-stage random cluster sample drawn from all Year 4 school classes in government and non-government schools in the five mainland Australian states of New South Wales, Victoria, South Australia, Western Australia and Queensland. The study was conducted between July 2003 and December 2004.

**Participants:** 1709 students from 88 schools (881 boys and 828 girls), representing 85% of the estimated target number of students. The class participation rate was 65%.

**Main outcome measures:** (i) Urinary iodine excretion (UIE) levels (compared with the criteria for the severity of iodine deficiency of the World Health Organization/International Council for the Control of Iodine Deficiency Disorders: iodine replete, UIE ≥ 100 µg/L; mild iodine deficiency, UIE 50–99 µg/L; moderate iodine deficiency, UIE 20–49 µg/L; severe iodine deficiency, UIE < 20 µg/L); (ii) Thyroid volumes measured by ultrasound (compared with new international reference values).

**Results:** Overall, children in mainland Australia are borderline iodine deficient, with a national median UIE of 104 µg/L. On a state basis, NSW and Victorian children are mildly iodine deficient, with median UIE levels of 89 µg/L and 73.5 µg/L, respectively. South Australian children are borderline iodine deficient, with a median UIE of 101 µg/L. Both Queensland and Western Australian children are iodine sufficient, with median UIE levels of 136.5 µg/L and 142.5 µg/L, respectively. Thyroid volumes in Australian schoolchildren are marginally increased compared with international normative data obtained from children living in iodine sufficient countries. There was no significant association between UIE and thyroid volume.

**Conclusion:** Our results confirm the existence of inadequate iodine intake in the Australian population, and we call for the urgent implementation of mandatory iodisation of all edible salt in Australia.

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### 1 Summary data on participating schoolchildren by state

State	Students participated/ students targeted	M:F ratio	Mean (SD) age (years)	Mean (SD) weight (kg)	Mean (SD) height (cm)	Mean (SD) body surface area (m <sup>2</sup> )	Median urinary iodine excretion (µg/L) (interquartile range)
NSW	427/400 (106%)	1:1	9.3±0.6	34.9±8.5	138.5±6.7	1.16±0.15	89.0 (65.0–123.5)
VIC	348/400 (87%)	1:0.8	9.7±0.5	38.2±8.9	141.0±6.8	1.22±0.15	73.5 (53.0–104.3)
SA	317/400 (79%)	1:0.9	9.0±0.5	35.3±7.9	137.3±7.3	1.16±0.15	101.0 (74.0–130.0)
WA	323/400 (80%)	1:0.8	8.9±0.6	32.8±7.6	136.9±6.4	1.11±0.14	142.5 (103.5–214.0)
QLD	294/400 (73%)	1:1.3	9.1±0.4	32.9±7.2	137.3±6.3	1.12±0.13	136.5 (104.3–183.8)
<b>Total</b>	<b>1709/2000 (85%)</b>	<b>1:0.9</b>	<b>9.2±0.6</b>	<b>34.9±8.3</b>	<b>138.3±6.9</b>	<b>1.20±0.10</b>	<b>104.0 (71.0–147.0)</b>

Research Ethics Committee and was approved and supported by the Departments of Health and the Departments of Education and Training in each state. Informed consent was obtained from parents or guardians of all participating school students.

#### Sample selection

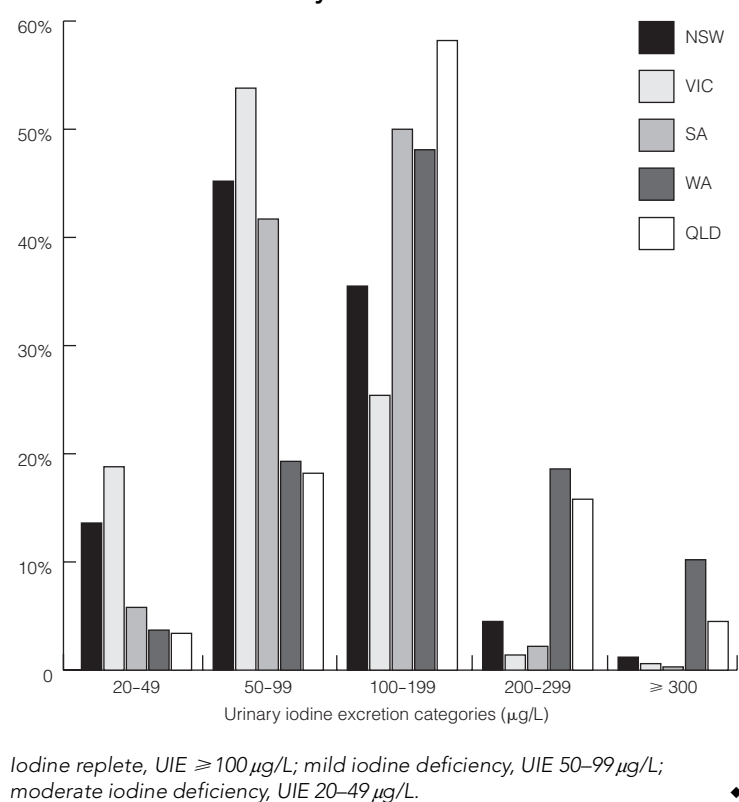
The sampling scheme was a one-stage random cluster sample, the cluster sampling frame comprising all Year 4 school classes in government and non-government schools. Children in Year 4 are typically aged 8–10 years. Cluster sampling is often used when a complete list of individuals in the population of interest is unavailable, but a complete list of groups or “clusters” of these individuals is available.<sup>11</sup>

Class sizes were obtained and entered into an MS Excel (Microsoft Corporation, Redmond, Calif, USA) database. If a school had more than one class in the target cluster population, each class was entered separately into the database. If the exact class size was unknown, the average class size of 25 was used. For composite classes, it was assumed that the children were distributed evenly between the two classes in the composite.

The Survey Toolbox software program Random Village component was used to generate a randomly sorted list of classes,<sup>11</sup> which were then selected in order until the target sample size (400 students from each state) was achieved.

If selected schools declined the invitation to participate in the study, more schools

### 2 Relative frequency distribution of urinary iodine excretion (UIE) levels in schoolchildren by state



were selected in the same fashion. Tasmania was excluded from the study because a voluntary iodine fortification program, using iodised salt in bread, is ongoing in that state; and the Northern Territory was excluded for logistical reasons.

#### Field studies

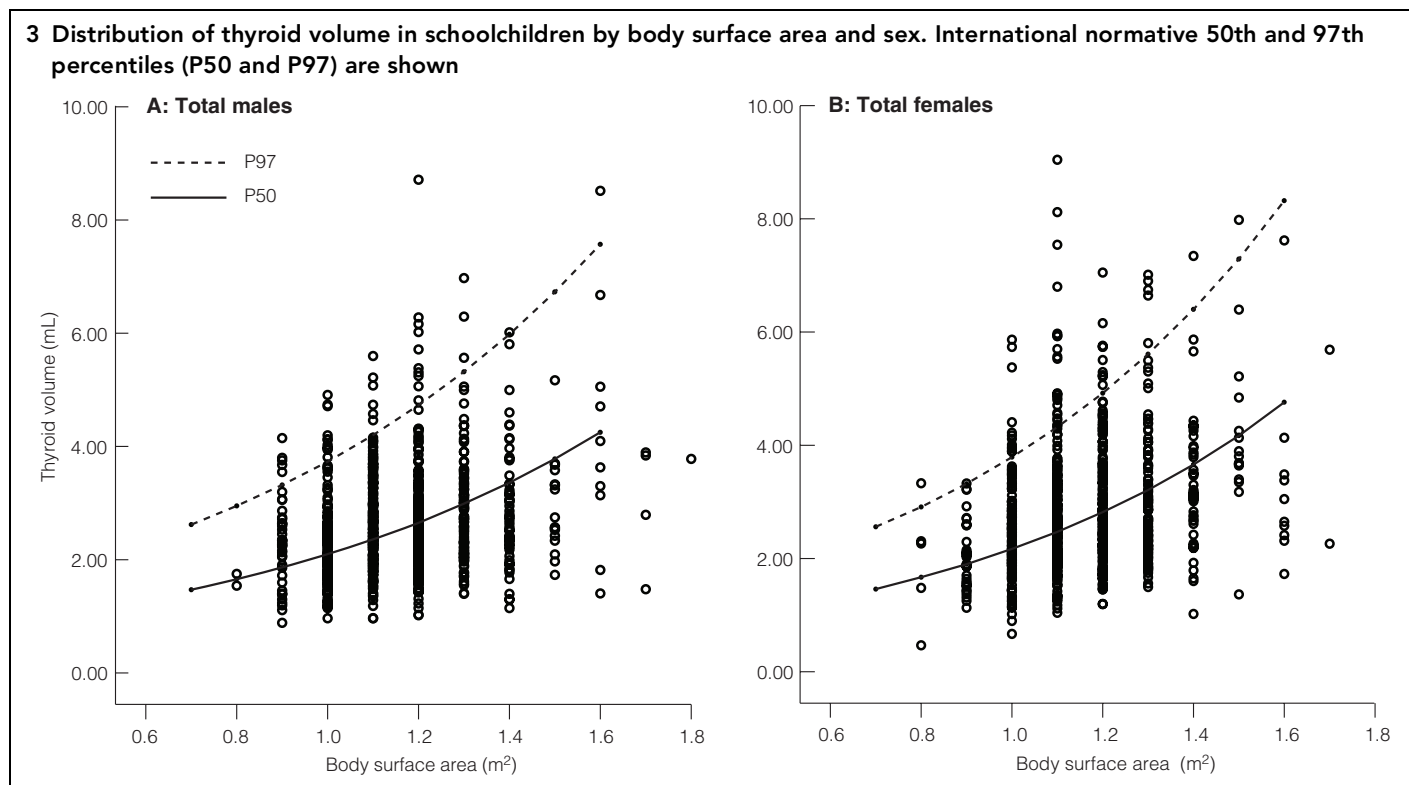
Our study, which aimed to provide a snapshot of iodine nutrition in Australian schoolchildren, aged 8–10 years, occurred between July 2003 and December 2004. We used a purpose-built vehicle known as a “ThyroMobil” (Merck, Darmstadt, Ger-

many), which has been used for similar studies of iodine nutrition in other countries.<sup>12–14</sup> At each school, students who brought their first morning urine sample to school with a pre-labelled specific identification number were registered, weighed and their height was measured (to calculate body surface area [BSA]) and they underwent a thyroid ultrasound scan.

#### Thyroid volume measured by ultrasound

Most of the thyroid volume measurements were performed with a Siemens Sonoline using a 7.5 MHz 6 cm linear transducer (Siemens, Munich, Germany). Thyroid ultrasound measurements were performed with the child in a supine position with the neck fully extended. For logistical reasons, there were three schools in South Australia and six schools in Western Australia where ultrasound measurement of thyroid volume was performed using a portable ultrasound machine (LOGIQ 100 PRO, GE Medical Systems, Mass, USA) calibrated against the Siemens machine in the ThyroMobil.

Ultrasound measurement of thyroid volume in children in NSW, South Australia, Western Australia and Queensland was performed either by the experienced reference sonographer (CJE) or by one other observer from each state, who undertook a calibration exercise with the reference sonographer to ensure uniformity of technique and measurement parameters. The between-observer variation was less than 10% in these states. In Victoria,



thyroid ultrasound measurements were performed by two independent investigators who had received instruction from the reference sonographer, but it was not possible at the time to undertake a calibration exercise in the field.

The volume of each thyroid lobe was calculated using the formula: volume = length  $\times$  width  $\times$  thickness  $\times$  0.479. Thyroid volume is the volume of the two lobes combined. Age- and BSA-specific 50th and 97th percentiles for thyroid volume were calculated for boys and girls. BSA was calculated by the formula: BSA = weight (kg)<sup>0.425</sup>  $\times$  height (cm)<sup>0.725</sup>  $\times$  71.84  $\times$  10<sup>-4</sup>.

If a child was found to have a pathologically enlarged thyroid gland during the survey, the abnormal finding was reported to parents or guardians of these children through their schools, with a general recommendation to seek further medical attention.

### Urinary iodine levels

Urine samples were collected and stored frozen in the ThyroMobil before being couriered back to Sydney where they were stored at  $-20^{\circ}\text{C}$  until assayed. Urinary iodine measurements were performed by ammonium persulfate digestion<sup>15</sup> before a Sandell-Kolthoff reaction in a microtitre plate format.<sup>16</sup>

### Statistical analysis

SPSS for Windows (version 12, SPSS Inc, Chicago, Ill, USA) was used to analyse the

data. Two-tailed tests with a significance level of 5% were used throughout. The thyroid volume and UIE data were log-transformed to approximate normality before analysis. General linear models were fitted to the log thyroid volume data and to the log UIE data to assess the joint effects of sex of the student and state, and of the covariates BSA or age. When a significant effect of state was detected, after adjusting for sex and BSA or age, pairwise multiple comparisons between states were corrected using the Bonferroni method. Spearman's rank correlation was used to quantify the association between thyroid volume and UIE.

## RESULTS

### Sample

Eighty-eight classes (in 88 schools), from the 135 classes selected across five states, participated in the study, with a class participation rate of 65%. In total, 1709 students (comprising 881 boys and 828 girls) participated, representing 85% of the estimated target number of students (Box 1).

### Urinary iodine excretion

Box 1 shows the median UIE levels and interquartile ranges for the five states, and in Box 2 the relative frequency distributions of

UIE level by state are grouped according to World Health Organization criteria for severity of iodine deficiency.<sup>17</sup> Box 2 also shows that, in NSW and Victoria, 58.8% and 72.6% of children, respectively, had UIE levels  $< 100\ \mu\text{g/L}$ . Similarly, close to half the South Australian children (47.5%) had UIE concentrations  $< 100\ \mu\text{g/L}$ , indicating inadequate iodine nutrition.

The log UIE levels varied markedly by state ( $P < 0.001$ ). After adjusting for sex, BSA and age, Victorian children had significantly lower adjusted levels than children in the other four states ( $P < 0.001$ ). There was no statistically significant difference between the adjusted log UIE levels for children from NSW and South Australia ( $P = 0.06$ ), which were significantly less ( $P < 0.001$ ) than those for children from Western Australia and Queensland, which were comparable ( $P = 0.8$ ).

### Thyroid volume

Box 3 shows the distribution of ultrasound measured thyroid volume (mL) by BSA and sex for children from all five states, and gives the international normative 50th and 97th percentile values (P50 and P97).<sup>18</sup> Box 4 shows the percentage of study children with a thyroid volume exceeding the P50 or P97 by sex, state and BSA or age. With the exception of Victoria, Australian children had relatively larger thyroid glands com-

#### 4 Percentage of children with a thyroid volume (mL) greater than the new international standard 50th and 97th percentile values (P50 and P97)

State	Based on body surface area (95% CI)		Based on age (95% CI)	
	Boys	Girls	Boys	Girls
<b>Percentage &gt; international standard P50</b>				
NSW	58.1 (51.3–64.9)	57.6 (50.9–64.3)	61.3 (54.6–68.0)	63.8 (57.3–70.3)
VIC	18.0 (12.6–23.5)	18.3 (12.2–24.4)	24.9 (18.8–31.0)	26.1 (19.2–33.1)
SA	41.0 (33.5–48.4)	46.3 (38.3–54.3)	50.9 (43.3–58.5)	54.4 (46.4–62.4)
WA	70.0 (63.1–76.9)	71.6 (64.4–78.9)	75.3 (68.8–81.8)	81.1 (74.8–87.4)
QLD	55.0 (46.5–63.6)	50.0 (42.3–57.5)	61.5 (53.2–69.9)	60.6 (53.1–68.2)
<b>Total</b>	<b>47.7 (44.3–51.0)</b>	<b>49.3 (45.8–52.7)</b>	<b>53.9 (50.6–57.3)</b>	<b>57.6 (54.2–60.9)</b>
<b>Percentage &gt; international standard P97</b>				
NSW	3.9 (1.3–6.6)	7.1 (3.7–10.6)	6.4 (3.0–9.7)	10.0 (5.9–14.1)
VIC	0	0	0	0.7 (0–1.9)
SA	4.8 (1.6–8.1)	10.1 (5.2–14.9)	6.6 (2.8–10.3)	10.7 (5.8–15.7)
WA	11.2 (6.4–15.9)	14.9 (9.1–20.6)	11.8 (6.9–16.6)	19.6 (13.2–26.0)
QLD	2.3 (0–4.9)	3.1 (0.4–5.8)	3.1 (0.1–6.0)	5.6 (2.1–9.2)
<b>Total</b>	<b>4.4 (3.0–5.8)</b>	<b>7.0 (5.2–8.7)</b>	<b>5.6 (4.0–7.1)</b>	<b>9.3 (7.3–11.3)</b>

pared with new international reference values for thyroid gland size in iodine-sufficient schoolchildren when adjusted by BSA or by age.

#### Adjusting for body surface area

In the general linear model of log thyroid volume, containing the main effects of sex and state and their interaction, and adjusting for BSA, the interaction term was not significant ( $P=0.8$ ). Sex and state were statistically significantly associated with thyroid volume after adjusting for BSA ( $P<0.001$  in each case). Overall, boys had smaller thyroid volumes after adjusting for BSA and state ( $P<0.001$ ). After adjusting for BSA and sex, Victorian children had significantly smaller thyroid volumes than those of the other four states ( $P<0.001$ ). Sex- and BSA-adjusted thyroid volumes were comparable for children from South Australia and Queensland ( $P=0.9$ ) who, in turn, had significantly smaller glands than NSW or Western Australian children ( $P<0.001$ ), who had the largest thyroid volumes.

#### Adjusting for age

In the general linear model of log thyroid volume, containing the main effects of sex and state and their interaction, and adjusting for age, the interaction term was not significant ( $P=0.8$ ). Sex and state were statistically significantly associated with thyroid volume after adjusting for age ( $P<0.001$  in each case). Overall, boys had smaller thyroid volumes after adjusting for age and state ( $P<0.001$ ). After adjusting for

age and sex, Victorian children had smaller measured thyroid volumes than children in the other four states ( $P<0.001$ ). Sex- and age-adjusted thyroid volumes were comparable for children from South Australia and Queensland ( $P=1.0$ ) who, in turn, had significantly smaller glands than NSW or Western Australian children ( $P<0.01$ ), whose gland sizes were comparable ( $P=0.1$ ).

#### Association between urinary iodine excretion and thyroid volume

There was no significant overall association between UIE and measured thyroid volume ( $r=0.02$ ,  $P=0.4$ ), although a very weak negative association was detected when the Victorian data were omitted ( $r=-0.058$ ,  $P=0.04$ ).

#### DISCUSSION

The National Iodine Nutrition Study is the largest study of its kind ever carried out in Australia, involving over 1700 students from 88 schools across five states. Random cluster sampling of classes generated a sample quickly and easily and gave each class an equal chance of being chosen, thereby providing an unbiased estimate of the overall population value.

Mainland Australian children as a group are borderline iodine deficient. However, there are significant and unexpected variations across the continent, with, overall, Western Australian and Queensland children being iodine replete and Victorian and

NSW children being mildly iodine deficient. The results from NSW and Victoria are very similar to those previously reported.<sup>5-7</sup> Tasmania was not included, but a study in 2001, before a bread fortification program was commenced, showed that the median UIE in Tasmanian schoolchildren was  $84\mu\text{g/L}$ , with 20% having levels under  $50\mu\text{g/L}$ .<sup>10</sup> These results for Tasmania are similar to our results for NSW and Victoria.

About half (46.3%) of all the students tested in mainland Australia had UIE levels in the range of mild (36.7%) to moderate (9.6%) iodine deficiency. Most of these children came from south-eastern Australia. Victoria (18.8%) and NSW (13.6%) had the highest percentage in the range of 20–49  $\mu\text{g/L}$ .

Children living in Western Australia and Queensland are clearly ingesting more iodine than their counterparts living elsewhere in Australia. We are not sure of the reasons for these regional variations, but the most likely explanations include:

- possible differences in the proportion of the population using iodised salt;
- variations in regional milk iodine content; and
- drinking water iodine levels.

These possibilities are currently being investigated. For example, the limited number of drinking water samples collected during the survey showed relatively high iodine levels in water and milk in northern and central Queensland, which could explain why the UIE levels indicated iodine sufficiency in this state.

Thyroid ultrasound, along with measurement of UIE levels, has been recommended by the WHO and the International Council for the Control of Iodine Deficiency Disorders (ICCIDD) for monitoring the sustained impact of iodine deficiency control programs through universal salt iodisation. Since their introduction in 1997, the recommended normative values for thyroid volume in children have been revised on two occasions. The validity of the 1997 WHO/ICCIDD recommended values were challenged by a number of studies, including surveys using a Thyro-Mobil.<sup>14</sup> It was later agreed that these reference values had significantly overestimated thyroid volumes in children.<sup>18</sup> Indeed, in the study in Tasmania in 2000, 5.3% (based on age) and 5.9% (based on BSA) of Tasmanian boys had increased thyroid volumes when compared with the 1997 international normative values. When compared with the 2001 corrected WHO/ICCIDD reference values,<sup>18</sup> the prevalence of raised thyroid volumes increased to 24.6% (based on age) and

22% (based on BSA).<sup>10</sup> A new set of international reference values for thyroid volume assessed by ultrasound examination, based on studies of children living in areas of long-term iodine sufficiency, were released in 2003.<sup>19</sup> The new reference values once again have lowered the reference volumes compared with the 2001 corrected values. The thyroid volume results in our study were compared with these new international reference values.

The thyroid volume results showed varying degrees of increased thyroid size across Australian states when compared with the international reference standards. Western Australian children had the largest glands, despite having the highest median UIE level of the five states. The explanation for this phenomenon is not clear. It may relate to factors other than iodine, such as dietary or environmental goitrogens. No association between spot urinary iodine level and thyroid volume assessed by ultrasound examination among schoolchildren has been reported.<sup>20</sup> With the exception of Victorian schoolchildren, we can conclude that thyroid volumes in Australian schoolchildren are marginally increased when compared with international normative data obtained from children in iodine sufficient countries, but this increase does not allow us to conclude that iodine deficiency is causing endemic goitre in Australia.

The Victorian thyroid volume data were not in keeping with the general data trends in the other four states. Most of the thyroid volumes measured in Victorian children were below the 50th percentile and none were above the 97th percentile of the international standards. We believe that this is a consequence of methodological error underestimating thyroid volume, and highlights the importance of training and standardisation of methods among observers to minimise the measurement error.<sup>20</sup> Victoria was the only state in which thyroid ultrasound measurements were performed by two independent investigators. While these two investigators received some instruction from the reference sonographer, the in-field calibration exercise performed in other states was not logistically possible at the time the data were collected.

It is alarming that mild iodine deficiency is extensive in children living in NSW,

Victoria, Tasmania and South Australia and no action has been taken by public health authorities to increase iodine intake in our population. It is reasonable to assume that pregnant women and breastfeeding mothers are also iodine deficient, putting the next generation of children born in this country at risk of the neuropsychological consequences of iodine deficiency.<sup>21</sup> The implementation of mandatory fortification of all edible salt for human consumption is long overdue in this country.

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## COMPETING INTERESTS

None identified.

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